

Form PTO-1449		U.S. Department of Commerce Patent and Trademark Office							Atty. Docket No. 48231-AZ-PCT-US JPW/AJM/MML	Serial No. Not Yet Known
		INFORMATION DISCLOSURE CITATION (Use several sheets if necessary)							Applicants: Greenwald and Levitan	
									Filing Date	Group 1653

U.S. PATENT DOCUMENTS												
Examiner Initial		Document Number					Date	Name		Class	Subclass	Filing Date if Appropriate
		6	3	7	6	2	3	9	4/23/03	Baumeister		
		6	0	8	7	1	5	3	7/11/00	Greenwald		
		5	8	4	0	5	4	0	11/24/98	St. George-Hyslop		

FOREIGN PATENT DOCUMENTS														
		Document Number					Date	Country		Class	Subclass	Translation		
													Yes	No
WO		9	7	1	1	9	5	6	3/3/97	PCT				X

OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.)										
		European Search Report, dated November 20, 2002;								
		PCT International Search Report, dated January 21, 1997;								
		PCT Written Opinion, dated July 15, 1997;								
		Bai C. et al. "SKP1 connects cell cycle regulators to the ubiquitin proteolysis machinery through a novel motif, the F-box," Cell 86:263-74 (1996);								
		Brenner S. "The genetics of <i>Caenorhabditis elegans</i> ," Genetics. (1974) 77(1):71-94;								
		Daigle I. and Li C. "apl-1, a <i>Caenorhabditis elegans</i> gene encoding a protein related to the human beta-amyloid protein precursor" Proc. Natl. Acad. Sci. U.S.A. (1993) 90(24):12045-9;								
		Database dbEST, National Center for Biotechnology Information, National Library of Medicine, GenBank Accession No. H19012 (1995);								
		Database EMBL Accession No: U35660 (1995) Levitan D., Greenwald I., "Caenorhabditis elegans membrane protein (sel-12) mRNA" XP002176178;								
		Ellisen L.W. et al. "TAN-1, the human homolog of the <i>Drosophila</i> Notch gene, is broken by chromosomal translocations in T lymphoblastic neoplasms," Cell 66:649-61 (1991);								
		Fire A. et al. "A modular set of lacZ fusion vectors for studying gene expression in <i>Caenorhabditis elegans</i> " Gene (1990) 93(2):189-98;								

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	Gallahan D. and Callahan R. "Mammary tumorigenesis in feral mice: identification of a new int locus in mouse mammary tumor virus (Czech II)-induced mammary tumors," <i>J. Virol.</i> 61:66-74 (1987);
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	Greenwald I. "Structure/function studies of lin-12/Notch proteins" <i>Curr. Opin. Genet. Dev.</i> (1994) 4(4):556-62;
	Hedgecock E.M. and Herman R.K. "The ncl-1 gene and genetic mosaics of <i>Caenorhabditis elegans</i> " <i>Genetics</i> (1995) 141(3):989-1006;
	Hedgecock E.M. et al. "Genetics of cell and axon migrations in <i>Caenorhabditis elegans</i> " <i>Development</i> (1987) 100(3):365-82;
	Hubbard J. et al. "Sel-10 negative regulator of lin-12 activity in <i>C. elegans</i> , encodes a member of the CDC4 family of proteins," <i>Genes Dev.</i> 11:3182-93 (1997);
	Kimble J. "Alteration in cell lineage following laser ablation of cells in the somatic gonad of <i>C. elegans</i> ," <i>Dev. Biol.</i> 87:286-300 (1981);
	King R.W. et al. "How proteolysis drives the cell cycle," <i>Science</i> 274:1652-58 (1996);
	Levitin D. and Greenwald I. "Facilitation of lin-12-mediated signalling by sel-12, a <i>Caenorhabditis elegans</i> S182 Alzheimer's disease gene" <i>Nature</i> (1995) 377(6547): 351-4;
	Levy-Lahad E. et al. "Genomic structure and expression of STM2, the chromosome 1 familial Alzheimer disease gene" <i>Genomics</i> (1996) 34(2):198-204;
	Levy-Lahad E. et al. "Candidate gene for the chromosome 1 familial Alzheimer's disease locus" <i>Science</i> (1995) 269(5226):973-7;
	L'Hernault S.W. and Arduengo P.M. "Mutation of a putative sperm membrane protein in <i>Caenorhabditis elegans</i> prevents sperm differentiation but not its associated meiotic divisions" <i>J. Cell. Biol.</i> (1992) 119(1):55-68;
	Li X. and Greenwald I. "HOP-1, a <i>Caenorhabditis elegans</i> presenilin, appears to be functionally redundant with SEL-12 presenilin and to facilitate LIN-12 and GLP-1 signaling" <i>Proc. Natl. Acad. Sci. U.S.A.</i> (1997) 94(22):12204-9;
	Mello C.C. et al. "Efficient gene transfer in <i>C. elegans</i> : extrachromosomal maintenance and integration of transforming sequences" <i>EMBO J.</i> (1991) 10(12):3959-70;
	Neer E.J. et al. "The ancient regulatory-protein family of WD-repeat proteins," <i>Nature</i> 371:297-300 (1994);
	Robbins J. et al. "Mouse mammary tumor gene int-3: a member of the Notch gene family transforms mammary epithelial cells," <i>J. Virol.</i> 66:2594-99 (1992);

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	Rogaev E. I. et al. "Familial Alzheimer's disease in kindreds with missense mutations in a gene on chromosome 1 related to the Alzheimer's disease type 3 gene" <i>Nature</i> (1995) 376(6543):775-8;
	Seydoux G. and Greenwald I. "Cell autonomy of lin-12 function in a cell fate decision in <i>C. elegans</i> ," <i>Cell</i> 57:1237-45 (1989);
	Shen J. et al. "Skeletal and CNS defects in presenilin-1-deficient mice," <i>Cell</i> 89:629-39 (1997);
	Sherrington R. et al. "Cloning of a gene bearing missense mutations in early-onset familial Alzheimer's disease" <i>Nature</i> (1995) 375(6534):754-60;
	Stratagene Cloning Systems Catalog, 1993, pages 27, 31, 32, and 313;
	Struhl G. et al., "Intrinsic activity of the Lin-12 and Notch intracellular domains in vivo" <i>Cell</i> (1993) 74(2):331-45;
	Sundaram M. and Greenwald I. "Genetic and phenotypic studies of hypomorphic lin-12 mutants in <i>Caenorhabditis elegans</i> " <i>Genetics</i> (1993) 135(3):755-63;
	Sundaram M. and Greenwald I. "Suppressors of a lin-12 hypomorph define genes that interact with both lin-12 and glp-1 in <i>Caenorhabditis elegans</i> " <i>Genetics</i> (1993) 135(3):765-83;
	Tuck S. and Greenwald I. "Lin-25, a gene required for vulval induction in <i>C. elegans</i> ," <i>Genes Dev.</i> 9:341-57 (1995);
	Wen C. et al. "spr-2, a suppressor of the egg-laying defect caused by loss of sel-12 presenilin in <i>Caenorhabditis elegans</i> , is a member of the SET protein subfamily" <i>Proc. Natl. Acad. Sci. U.S.A.</i> (2000) 97(26):14524-9;
	Wilkinson H.A. and Greenwald I. "Spatial and temporal patterns of lin-12 expression during <i>C. elegans</i> hermaphrodite development" <i>Genetics</i> (1995) 141(2):513-26;
	Wilkinson H.A. et al. "Reciprocal changes in expression of the receptor lin-12 and its ligand lag-2 prior to commitment in a <i>C. elegans</i> cell fate decision" <i>Cell</i> (1994) 79(7):1187-98;
	Wilson R. et al. "2.2 Mb of contiguous nucleotide sequence from chromosome III of <i>C. elegans</i> ," <i>Nature</i> 368:32-38 (1994);
	Wong P.C. et al. "Presenilin-1 is required for Notch1 and DII 1 expression in the paraxial mesoderm," <i>Nature</i> 387:288-91 (1997); and
	Yochim J. and Byers B. "Structural comparison of the yeast cell division cycle gene CDC4 and a related pseudogene," <i>J. Mol. Biol.</i> 195:233-45 (1987).
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Sequence Listing

Applicants submit herewith (a) a paper copy of the Sequence Listing, inserted into the specification following the Abstract of the Disclosure, (b) a request to use the computer readable form (CRF) of the sequence listing submitted in prior application U.S. Serial No. 09/043,944, attached hereto as **Exhibit A**, and (c) a statement in accordance with 37 C.F.R. §1.821(f) attached hereto as **Exhibit B**, certifying that the CRF and written sequence listing contain the same sequence information.

Information Disclosure

In accordance with their duty of disclosure under 37 C.F.R. §1.56, applicants request that the following disclosures be made of record in the above-identified application pursuant to 37 C.F.R. §1.97(b). These references are also listed on the Form PTO-1449 attached hereto as **Exhibit C**. Copies of these references were submitted in connection with prior application U.S. Serial No. 09/043,944.

1. U.S. Patent No. 6,376,239, issued April 23, 2002, Baumeister;
2. U.S. Patent No. 6,087,153, issued July 11, 2000, Greenwald et al.;
3. U.S. Patent No. 5,840,540, issued November 24, 1998, Peter H. St. George-Hyslop et al.;
4. PCT International Application No. WO 97/11956, published April 3, 1997;
5. European Search Report, dated November 20, 2002;
6. PCT International Search Report, dated January 21, 1997;
7. PCT Written Opinion, dated July 15, 1997;

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8. Bai C. et al. "SKP1 connects cell cycle regulators to the ubiquitin proteolysis machinery through a novel motif, the F-box," *Cell* 86:263-74 (1996);
9. Brenner S. "The genetics of *Caenorhabditis elegans*," *Genetics*. (1974) 77(1):71-94;
10. Daigle I. and Li C. "apl-1, a *Caenorhabditis elegans* gene encoding a protein related to the human beta-amyloid protein precursor" *Proc. Natl. Acad. Sci. U.S.A.* (1993) 90(24):12045-9;
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13. Ellisen L.W. et al. "TAN-1, the human homolog of the *Drosophila* Notch gene, is broken by chromosomal translocations in T lymphoblastic neoplasms," *Cell* 66:649-61 (1991);
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37. Sherrington R. et al. "Cloning of a gene bearing missense mutations in early-onset familial Alzheimer's disease" *Nature* (1995) 375(6534):754-60;
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No fee, other than the enclosed filing fee, is deemed necessary in connection with the filing of this Preliminary Amendment. However, if any additional fee is required, authorization is hereby given to charge the amount of such fee to Deposit Account No. 03-3125.

If a telephone interview would be of assistance in advancing prosecution of the subject application, applicants' undersigned attorneys invite the Examiner to telephone them at the number provided below.

Respectfully submitted,

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